Amendments to the Claims

This listing of claims will replace all prior versions, and listings, of claims in the application:

1. (Currently Amended) <u>A</u> [[M]]method for in vitro detection of acute generalized inflammatory conditions (SIRS), comprising:

characterized in that

it comprises the following steps:

Isolation of isolating sample RNA from a sample of a mammal;

[[L]]]abelling of the sample RNA and/or at least one DNA being a gene or gene fragment specific for SIRS, with a detectable label.

[[C]]contacting the sample RNA with the DNA under hybridization conditions;

[[C]]contacting sample RNA representing a control for non-pathologic conditions, with at least one DNA, under hybridization conditions, whereby the DNA is a gene or gene fragment specific for SIRS;

[[Q]]quantitative detection of the label signals of the hybridized sample RNA and control RNA; and

[[C]]comparing the quantitative data of the label signals in order to determine whether the genes or gene fragments specific for SIRS are more expressed in the sample than in the control, or less.

 (Currently Amended) A [[M]]method for in vitro detection of sepsis and/or sepsis-like conditions.

characterized in that

Page 3 of 13

it comprises the following steps:

Isolation isolating of sample RNA from a sample of a mammal;

[[L]]labelling of the sample RNA and/or at least one DNA being a gene or gene fragment

specific for sepsis, with a detectable label.

[[C]]contacting the sample RNA with the DNA under hybridization conditions;

[[C]]contacting sample RNA representing a control for non-pathologic conditions, with at least one DNA, under hybridization conditions, whereby the DNA is a gene or gene fragment

specific for sepsis and/or sepsis-like conditions;

[[Q]]quantitative detection of the label signals of the hybridized sample RNA and control

RNA; and

[[C]]comparing the quantitative data of the label signals in order to determine whether

the genes or gene fragments specific for sepsis and/or sepsis-like conditions are more expressed in

the sample than in the control, or less.

3. (Currently Amended) A [[M]]method for in vitro detection of severe sepsis, comprising:

characterized in that

it comprises the following steps:

Isolation isolating of sample RNA from a sample of a mammal;

[[L]]labelling of the sample RNA and/or at least one DNA being a gene or gene fragment

specific for severe sepsis, with a detectable label.

[[C]]contacting the sample RNA with the DNA under hybridization conditions;

{WP407935:1}

Docket No. 3535-027

Application No. 10/551,874 Second Preliminary Amendment Page 4 of 13

[[C]]ontacting sample RNA representing a control for non-pathologic conditions, with at least one DNA, under hybridization conditions, whereby the DNA is a gene or gene fragment specific for severe sensis;

[[Q]]quantitative detection of the label signals of the hybridized sample RNA and control RNA; and

[[C]]comparing the quantitative data of the label signals in order to determine whether the genes or gene fragments specific for severe sepsis are more expressed in the sample than in the control or less.

- (Currently Amended) The [[M]]method according to one of claim[[s]] 1 [[to 3]], characterized in that the control RNA is hybridized with the DNA before the measurement of the sample RNA and the label signals of the control RNA/DNA-complex is gathered and, if necessary, recorded in form of a calibration curve or table.
- (Currently Amended) The [[M]]method aecerding to one of claim[[s]] 1 [[to 4]], characterized in that unchanged genes from sample and/or control RNA are used as reference genes for the quantification.
- (Currently Amended) The [[M]]method according to one of claim[[s]] 1 [[to 5]], characterized in that mRNA is used as sample RNA.
- (Currently Amended) <u>The</u> [[M]]method according to ene of claim[[s]] 1 [[to 6]], characterized in that the DNA is arranged, particularly immobilized, on predetermined areas on a carrier in the form of a microarray.
- 8. (Currently Amended) The [[M]]method according to one of claim[[s]] 1 [[to 7]], characterized in that the method for early detection by means of differential diagnostics, for control of the clinical and therapeutic progress, for the individual risk evaluation in patients, for the evaluation whether the patient will respond to a specific treatment, as well as for post mortem diagnosis of SIRS and/or sepsis and/or severe sepsis and/or systemic infections and/or septic conditions and/or infections.

Application No. 10/551,874 Second Preliminary Amendment Page 5 of 13

- (Currently Amended) The [[M]]method according to one of claim[[s]] 1 [[to 8]], characterized in that the sample is selected from the following group: body fluids, in particular blood, liquor, urine, ascitic fluid, seminal fluid, saliva, puncture fluid, cell content, or a mixture thereof.
- (Currently Amended) The [[M]]method aecording to one of claim[[s]] 1 [[to 9]], characterized in that cell samples are subjected a lytic treatment, if necessary, in order to free their cell contents.
- (Currently Amended) The [[M]]method according to one of claim[[s]] 1 [[to 10]], characterized in that the mammal is a human.
- 12. (Currently Amended) The [[M]]method according to one of claim[[s]] 1 or 4 to 11, characterized in that the gene or gene segment specific for SIRS is selected from the group consisting of SEQUENCE ID No. III.1 to SEQUECE ID No. III.4168, as well as gene fragments thereof with 5-2000 or more, preferably 20-200, more preferably 20-80 nucleotides.
- 13. (Currently Amended) The [[M]]method aecerding to one of claim[[s]] 2 or 4 to 11, characterized in that the gene or gene segment specific for sepsis and/or sepsis-like conditions is selected from the group consisting of SEQUENCE ID No. I.1 to SEQUECE ID No. I.6242, as well as gene fragments thereof with 5-2000 or more, preferably 20-200, more preferably 20-80 nucleotides.
- 14. (Currently Amended) The [[M]]method according to one of claim[[s]] 3 or 4 to -11, characterized in that the gene or gene segment specific for severe sepsis is selected from the group consisting of SEQUENCE ID No. II.1 to SEQUECE ID No. II.130, as well as gene fragments thereof with 5-2000 or more preferably 20-200, more preferably 20-80 nucleotides.
- (Currently Amended) <u>The [[M]]method aecording to one</u> of claim[[s]] 1 [[to 14]], characterized in that at least 2 to 100 different cDNAs are used.
- (Currently Amended) <u>The [[M]]method according to one</u> of claim[[s]] 1 [[to 15]], characterized in that at least 200 different cDNAs are used.

- (Currently Amended) <u>The [[M]]method according to one of claim[[s]] 1 [[to 16]], characterized in that at least 200 to 500 different cDNAs are used.</u>
- (Currently Amended) <u>The [[M]]method aecording to one</u> of claim[[s]] 1 [[to 17]], characterized in that at least 500 to 1000 different cDNAs are used.
- (Currently Amended) The [[M]]method according to one of claim[[s]] 1 [[to 18]], characterized in that at least 1000 to 2000 different cDNAs are used.
- (Currently Amended) The [[M]]method according to one of claim[[s]] 1 [[to 19]], characterized in that the cDNA of the genes listed in claims 12, 13 und 14 is SEQUENCE ID No. III.1 to SEQUECE ID No. III.4168, SEQUENCE ID No. I.1 to SEQUECE ID No. I.6242 and SEQUENCE ID No. II.1 to SEQUECE ID No. II.130 replaced by synthetic analoga as well as peptidonucleic acids.
- (Currently Amended) The [[M]]method according to of claim 20, characterized in that the
 synthetic analoga of the listed genes comprise 5-100, in particular approximately 70, base pairs.
- (Currently Amended) The [[M]]method aecording to one of claim[[s]] 1 [[to 21]], characterized in that a radioactive label, in particular ³²P, ¹⁴C, ¹²⁵I, [[¹⁵⁵Ep]] ¹⁵⁵Eu, ³³P or ³H is used as detectable label.
- 23. (Currently Amended) The [[M]]method according to one of claim[[s]] 1 [[to 22]], characterized in that a non-radioactive label is used as detectable label, in particular a color- or fluorescence label, an enzyme label or immune label, and/or quantum dots or an electrically measurable signal, in particular the change in potential, and/or conductivity and/or capacity by hybridizations.
- (Currently Amended) The [[M]]method according to one of claim[[s]] 1 [[to 23]], characterized in that the sample RNA and control RNA bear the same label.
- (Currently Amended) The [[M]]method according to one of claim[[s]] 1 [[to 24]], characterized in that the sample RNA and control RNA bear different labels.

Application No. 10/551,874 Second Preliminary Amendment Page 7 of 13

- (Currently Amended) The [[M]]method according to one of claim[[s]] 1 [[to 25]], characterized in that the immobilized probes bear a label.
- (Currently Amended) The [[M]]method according to one of claim[[s]] 1 [[to 26]], characterized in that the cDNA probes are immobilized on glass or plastics.
- (Currently Amended) The [[M]]method according to one of claim[[s]] 1 [[to 27]],
 characterized in that the individual cDNA molecules are immobilized on the carrier material by
 means of a covalent binding.
- 29. (Currently Amended) The [[M]]method aecording to one of claim[[s]] 1 [[to 28]], characterized in that the individual cDNA molecules are immobilized onto the carrier material by means of adsorption, in particular by means of electrostatic and/or dipole-dipole and/or hydrophobic interactions and/or hydrogen bridges.
- 30. (Currently Amended) A [[M]]method for in vitro detection of SIRS, comprising:

characterized in that

it comprises the following steps:

Isolation of isolating sample peptides from a sample of a mammal;

[[L]]labelling of the sample peptides with a detectable label;

[[C]]contacting the labelled sample peptides with at least one antibody or its binding fragment, whereby the antibody binds a peptide or peptide fragment specific for SIRS;

[[C]]contacting the labelled control peptides originating from healthy subjects, with at least one antibody or its binding fragment immobilized on a carrier in form of a microarray, whereby the antibody binds a peptide or peptide fragment specific for SIRS;

[[Q]]quantitative detection of the label signals of the sample peptides and the control peptides;

Docket No. 3535-027

Application No. 10/551,874 Second Preliminary Amendment

Page 8 of 13

[[C]]comparing the quantitative data of the label signals in order determine whether the

genes or gene fragments specific for SIRS are more expressed in the sample than in the control,

or less.

31. (Currently Amended) A [[M]]method for in vitro detection of sepsis and/or sepsis-like

conditions, comprising:

characterized in that

it comprises the following steps:

Isolation of isolating sample peptides from a sample of a mammal;

[[L]]labelling of the sample peptides with a detectable label;

[[C]]contacting the labelled sample peptides with at least one antibody or its binding

fragment, whereby the antibody binds a peptide or peptide fragment specific for sepsis and/or

sepsis-like conditions;

[[C]]contacting the labelled control peptides stemming from healthy subjects, with at least

one antibody or its binding fragment immobilized on a carrier in form of a microarray, whereby the antibody binds a peptide or peptide fragment specific for sepsis and/or sepsis-like conditions;

[[O]]quantitative detection of the label signals of the sample peptides and the control

peptides; and

[[C]]comparing the quantitative data of the label signals in order to be able to determine

whether the genes or gene fragments specific for sepsis and/or sepsis-like conditions are more

expressed in the sample than in the control, or less.

32. (Currently Amended) A [[M]]method for in vitro detection of severe sepsis, comprising:

characterized in that

it comprises the following steps:

{WP407935:1}

Application No. 10/551,874
Second Preliminary Amendment
Page 9 of 13

Isolation of isolating sample peptides from a sample of a mammal;

[[L]]labelling of the sample peptides with a detectable label;

[[C]]contacting the labelled sample peptides with at least one antibody or its binding fragment, whereby the antibody binds a peptide or peptide fragment specific for severe sepsis;

[[C]]contacting the labelled control peptides originating from healthy subjects, with at least one antibody or its binding fragment immobilized on a carrier in form of a microarray, whereby the antibody binds a peptide or peptide fragment specific for severe sepsis;

[[Q]]quantitative detection of the label signals of the sample peptides and the control peptides; and

[[C]]comparing the quantitative data of the label signals in order to determine whether the genes or gene fragments specific for severe sepsis are more expressed in the sample than in the control, or less.

- (Currently Amended) The [[M]]method according to one of claim[[s]] 30 [[to 32]], characterized in that the antibody is immobilized on an array in form of a microarray.
- (Currently Amended) The [[M]]method according to one of claim[[s]] 30 [[to 33]], characterized in that it is formed as immunoassay.
- 35. (Currently Amended) The [[M]]method according to one of claim[[s]] 30 [[to 34]], characterized in that the method is used for early detection by means of differential diagnostics, for control of the clinic and therapeutic progress, for risk evaluation for patients as well as for post mortem diagnosis of SIRS and/or sepsis and/or severe sepsis and/or systemic infections and/or septic conditions and/or infections.
- 36. (Currently Amended) The [[M]]method according to one of claim[[s]] 30 [[to 35]], characterized in that the sample is selected from the following group: body fluids, in particular blood, liquor, urine, ascitic fluid, seminal fluid, saliva, puncture fluid, cell content, or a mixture thereof

Application No. 10/551,874 Second Preliminary Amendment Page 10 of 13

- (Currently Amended) The [[M]]method aecording to one of claim[[s]] 30 [[to 36]],
 characterized in that cell samples are subjected a lytic treatment, if necessary, in order to free
 their cell contents.
- (Currently Amended) The [[M]]method according to one of claim[[s]] 30 [[to 37]], characterized in that the mammal is a human.
- 39. (Currently Amended) The [[M]]method according to one of claim[[s]] 30 or 33 to 38, characterized in that the peptide specific for SIRS is an expression product of a gene or gene fragment selected from the group consisting of SEQUENCE ID No. III.1 to SEQUECE ID No. III.4168, as well as gene fragments thereof with 5-2000 nucleotides or more, preferably 20-200, more preferable 20-80 nucleotides.
- 40. (Currently Amended) The [[M]]method according to one of claim[[s]] 31 or 33 to 38, characterized in that the peptide specific for sepsis and/or sepsis-like conditions is an expression product of a gene or gene fragment selected from the group consisting of SEQUENCE ID No. I.1 to SEQUECE ID No. I.6242, as well as gene fragments thereof with 5-2000 nucleotides or more, preferably 20-200, more preferable 20-80 nucleotides.
- 41. (Currently Amended) The [[M]]method according to one of claim[[s]] 32 or 33 to 38, characterized in that the peptide specific for severe sepsis is an expression product of a gene or gene fragment selected from the group consisting of SEQUENCE ID No. II.1 to SEQUECE ID No. II.130, as well as gene fragments thereof with 5-2000 or more, preferably 20-200, more preferably 20-80 nucleotides.
- (Currently Amended) The [[M]]method according to one of claim[[s]] 30 [[to 41]], characterized in that at least 2 to 100 different peptides are used.
- (Currently Amended) The [[M]]method according to one of claim[[s]] 30 [[to 42]], characterized in that at least 200 different peptides are used.
- (Currently Amended) The [[M]]method according to one of claim[[s]] 30 [[to 43]], characterized in that at least 200 to 500 different peptides are used.

- (Currently Amended) The [[M]]method according to one of claim[[s]] 30 [[to 44]], characterized in that at least 500 to 1000 different peptides are used.
- (Currently Amended) The [[M]]method according to one of claim[[s]] 30 to 45, characterized
 in that at least 1000 to 2000 different peptides are used.
- (Currently Amended) The [[M]]method according to one of claim[[s]] 30 [[to 46]], characterized in that a radioactive label, in particular ³²P, ¹⁴C, ¹²⁵I, [[¹⁵⁵Ep]] ¹⁵⁵Eu, ³³P or ³H is used as detectable label.
- 48. (Currently Amended) The [[M]]method according to one of claim[[s]] 30 [[to 47]], characterized in that a non-radioactive label is used as detectable label, in particular a color- or fluorescence label, an enzyme label or immune label, and/or quantum dots or an electrically measurable signal, in particular the change in potential, and/or conductivity and/or capacity by hybridizations.
- (Currently Amended) The [[M]]method aecording to one of claim[[s]] 30 [[to 48]], characterized in that the sample peptides and control peptides bear the same label.
- (Currently Amended) The [[M]]method according to one of claim[[s]] 30 [[to 49]], characterized in that the sample peptides and control peptides bear different labels.
- 51. (Currently Amended) The [[M]]method according to—one of claim[[s]] 30 [[to 50]], characterized in that the probes used are peptides to which labelled antibodies are bound, which cause a change of signal of the labelled antibodies by change of conformation when binding to the sample peptides.
- (Currently Amended) The [[M]]method according to one of claim[[s]] 30 [[to 51]], characterized in that the peptide probes are immobilized on glass or plastics.
- (Currently Amended) The [[M]]method according to one of claim[[s]] 30 [[to 52]],
 characterized in that the individual peptide molecules are immobilized onto the carrier material
 by means of a covalent binding.

Docket No. 3535-027

Application No. 10/551,874 Second Preliminary Amendment Page 12 of 13

- 54. (Currently Amended) The [[M]]method aecerding to one of claim[[s]] 30 [[to 53]], characterized in that the individual peptide molecules are immobilized on the carrier material by means of adsorption, in particular by means of electrostatic and/or dipole-dipole and/or hydrophobic interactions and/or hydrogen bridges.
- 55. (Currently Amended) The [[M]]method aecerding to one of claim[[s]] 30 [[to 54]], characterized in that the individual peptide molecules are detected by means of monoclonal antibodies or their binding fragments.
- (Currently Amended) The [[M]]method aecording to—one of claim[[s]] 30 [[to 55]], characterized in that the determination of individual peptides by means of immunoassay or precipitation assay is carried out using monoclonal antibodies.
- 57. (Cancelled)
- 58. (Cancelled)
- (Cancelled)